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Shirley,

5/27/06

Comments on the FAOMs for 10/743,354:

1. I see that the only claim under examination is a composition claim. Why then does there need to be a step in such a claim as you seem to require via the 112, 2nd paragraph, rejection on page 2 of the action? ~~deleted -~~

2. On the bottom of page 3 of the action the "similar compound" reasoning is clearly insufficient to provide a prima facie case for rejecting the elected invention under 103. We have discussed this improper "similar compound" reasoning as being improper.

Reviewingly,



Ardin

DETAILED ACTION

The response to the restriction requirement filed **February 27, 2006** presents election of group 1 claim 1 without traverse is acknowledged.

Status of claims

Claims 2-5 are with drawn and claim 1 is pending in this office action.

Information Disclosure Statement

The information disclosure statement filed December 22, 2003 has been considered however items 45 and 41 of page 1 are not considered because a search report is not a published document. It has been placed in the application file, but the information referred to therein has not been considered as to the merits.

Claim Rejections - 35 USC § 112

Claim 1 is rejected under 35 U.S.C. 112, second paragraph, as being incomplete for omitting essential steps, such omission amounting to a gap between the steps. See MPEP § 2172.01. The omitted steps are: the utility of the said pharmaceutical composition. What is the pharmaceutical composition used for? The limitation should be set forth in the claim. Examiner suggest restating claim 1 to include a method step from the withdrawn claims "a pharmaceutical composition for treating $\alpha\text{v}\beta_3$.

Claim Rejections - 35 USC § 103

This application currently names joint inventors. In considering patentability of the claims under 35 U.S.C. 103(a), the examiner presumes that the subject matter of

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the various claims was commonly owned at the time any inventions covered therein were made absent any evidence to the contrary. Applicant is advised of the obligation under 37 CFR 1.56 to point out the inventor and invention dates of each claim that was not commonly owned at the time a later invention was made in order for the examiner to consider the applicability of 35 U.S.C. 103(c) and potential 35 U.S.C. 102(e), (f) or (g) prior art under 35 U.S.C. 103(a).

Claim 1 is rejected under 35 U.S.C. 103(a) as being unpatentable over Duggan et al WO 99/30709 in view of Nagarajan et al. US 6,933,304 B2.

Duggan et al. teach the synthesis of various heteroarylalkenoic acids (see pages 68-144).

Nagarajan et al. teach the synthesis of various heteroarylalkenoic acids (see cols. 26-154).

The instant invention differs from that disclosed by the above reference in that the particular compound 3-(3-tert-butyl-5-iodophenyl)-4-{3-[3-(5,6,7,8-tetrahydro-1,8-naphthyridin-2-yl)propyl]-1,2,4-oxadiazol-5-yl}butanoic acid in the instant claim versus

3-(4-fluorophenyl)-4-{3-[3-(5,6,7,8-tetrahydro-1,8-naphthyridin-2-yl)propyl]-1,2,4-oxadiazol-5-yl}butanoic acid trifluoroacetate

in the cited reference of Nagarajan et al..

Both cited references teach the synthesis of heteroarylalkenoic acids that is similar to the above mentioned compound. One of ordinary skill in the art would be motivated to employ the above teachings and synthesis compounds to that of the claimed invention because, Nagarajan et al. teach the general synthetic process of making such compounds in a pharmaceutical composition.

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One of ordinary skill in the art would have recognize the modification of the parent drug to yield a pro drug because the Nagarajan et al. with that of Duggan et al. teach the general synthethic means of making the compound of same structural

The general synthetic sequences for preparing the compounds useful in the present invention are outlined in Schemes 1-12. Both an explanation of, and the actual procedures for, the various aspects of the present invention are described where appropriate. The following Schemes and Examples are intended to be merely illustrative of the present invention, and not limiting thereof in either scope or spirit. Those with skill in the art will readily understand that known variations of the conditions and processes described in the Schemes and Examples can be used to synthesize the compounds of the present invention.

similarity with different moieties.

Thus, the claimed invention was prima facia obvious to make and use at the time it was made.

Any inquiry concerning this communication or earlier communications from the examiner should be directed to Shirley V. Gembah whose telephone number is 571-272-8504. The examiner can normally be reached on 8:30 -5:00, Monday- Friday.

If attempts to reach the examiner by telephone are unsuccessful, the examiner's supervisor, Ardin Marschel can be reached on 571-272-0718. The fax phone number for the organization where this application or proceeding is assigned is 571-273-8300.

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Information regarding the status of an application may be obtained from the Patent Application Information Retrieval (PAIR) system. Status information for published applications may be obtained from either Private PAIR or Public PAIR. Status information for unpublished applications is available through Private PAIR only. For more information about the PAIR system, see <http://pair-direct.uspto.gov>. Should you have questions on access to the Private PAIR system, contact the Electronic Business Center (EBC) at 866-217-9197 (toll-free).

SVG
5/10/06

3-(3-Furyl)-4-{3-[3-(5,6,7,8-tetrahydro-1,8-naphthyridin-2-yl)propyl]-1,2,4-oxadiazol-5-yl}butanoic acid trifluoroacetate;

35 4-{3-[3-(5,6,7,8-Tetrahydro-1,8-naphthyridin-2-yl)propyl]-1,2,4-oxadiazol-5-yl}-3-thien-3-ylbutanoic acid trifluoroacetate;

3-(2,3-Dihydro-1,4-benzodioxin-6-yl)-4-{3-[3-(5,6,7,8-tetrahydro-1,8-naphthyridin-2-yl)propyl]-1,2,4-oxadiazol-5-yl}butanoic acid hydrochloride;

40 4-{3-[3-(5,6,7,8-Tetrahydro-1,8-naphthyridin-2-yl)propyl]-1,2,4-oxadiazol-5-yl}-3-[3-(trifluoromethoxy)phenyl]butanoic acid hydrochloride;

4-{3-[3-(5,6,7,8-Tetrahydro-1,8-naphthyridin-2-yl)propyl]-1,2,4-oxadiazol-5-yl}-3-(3,4,5-trifluorophenyl)butanoic acid hydrochloride;

3-(2,2-Difluoro-1,3-benzodioxol-5-yl)-4-{3-[3-(5,6,7,8-tetrahydro-1,8-naphthyridin-2-yl)propyl]-1,2,4-oxadiazol-5-yl}butanoic acid hydrochloride;

45 3-[3-Fluoro-5-(trifluoromethyl)phenyl]-4-{3-[3-(5,6,7,8-tetrahydro-1,8-naphthyridin-2-yl)propyl]-1,2,4-oxadiazol-5-yl}butanoic acid hydrochloride;

3-(6-Methoxy-2-naphthyl)-4-{3-[3-(5,6,7,8-tetrahydro-1,8-naphthyridin-2-yl)propyl]-1,2,4-oxadiazol-5-yl}butanoic acid hydrochloride;

50 3-(6-Methoxypyridin-3-yl)-4-{3-[3-(5,6,7,8-tetrahydro-1,8-naphthyridin-2-yl)propyl]-1,2,4-oxadiazol-5-yl}butanoic acid;

3-(4-Cyanophenyl)-4-{3-[3-(5,6,7,8-tetrahydro-1,8-naphthyridin-2-yl)propyl]-1,2,4-oxadiazol-5-yl}butanoic acid;

3-(3-Cyanophenyl)-4-{3-[3-(5,6,7,8-tetrahydro-1,8-naphthyridin-2-yl)propyl]-1,2,4-oxadiazol-5-yl}butanoic acid;

55 3-benzyl-4-{3-[3-(5,6,7,8-tetrahydro-1,8-naphthyridin-2-yl)propyl]-1,2,4-oxadiazol-5-yl}butanoic acid trifluoroacetate;

3-(4-fluoro-3-methoxyphenyl)-4-{3-[3-(5,6,7,8-tetrahydro-1,8-naphthyridin-2-yl)propyl]-1,2,4-oxadiazol-5-yl}butanoic acid trifluoroacetate;

60 3-(3-Fluoro-5-methoxyphenyl)-4-{3-[3-(5,6,7,8-tetrahydro-1,8-naphthyridin-2-yl)propyl]-1,2,4-oxadiazol-5-yl}butanoic acid hydrochloride;

3-(2-Methyl-1,3-benzothiazol-5-yl)-4-{3-[3-(5,6,7,8-tetrahydro-1,8-naphthyridin-2-yl)propyl]-1,2,4-oxadiazol-5-yl}butanoic acid hydrochloride;

3-[2-(4-Chlorophenyl)-1,3-thiazol-5-yl]-4-{3-[3-(5,6,7,8-tetrahydro-1,8-naphthyridin-2-yl)propyl]-1,2,4-oxadiazol-5-yl}butanoic acid hydrochloride;

65 3-[2-(4-Methoxyphenyl)-1,3-thiazol-5-yl]-4-{3-[3-(5,6,7,8-tetrahydro-1,8-naphthyridin-2-yl)propyl]-1,2,4-oxadiazol-5-yl}butanoic acid hydrochloride;

- 3-(2-Methyl-1,3-benzothiazol-5-yl)-4-{3-[3-(5,6,7,8-tetrahydro-1,8-naphthyridin-2-yl)propyl]-1,2,4-oxadiazol-5-yl}butanoic acid hydrochloride;
- 70 3-[2-(4-Fluorophenyl)-1,3-thiazol-5-yl]-4-{3-[3-(5,6,7,8-tetrahydro-1,8-naphthyridin-2-yl)propyl]-1,2,4-oxadiazol-5-yl}butanoic acid hydrochloride;
- 3-[2-(3,5-Difluorophenyl)-1,3-thiazol-5-yl]-4-{3-[3-(5,6,7,8-tetrahydro-1,8-naphthyridin-2-yl)propyl]-1,2,4-oxadiazol-5-yl}butanoic acid hydrochloride;
- 3-[2-(3,4-Difluorophenyl)-1,3-thiazol-5-yl]-4-{3-[3-(5,6,7,8-tetrahydro-1,8-naphthyridin-2-yl)propyl]-1,2,4-oxadiazol-5-yl}butanoic acid hydrochloride;
- 75 3-[2-(2-Furyl)-1,3-thiazol-5-yl]-4-{3-[3-(5,6,7,8-tetrahydro-1,8-naphthyridin-2-yl)propyl]-1,2,4-oxadiazol-5-yl}butanoic acid hydrochloride;
- 3-(3,4-Dimethoxyphenyl)-4-{3-[3-(5,6,7,8-tetrahydro-1,8-naphthyridin-2-yl)propyl]-1,2,4-oxadiazol-5-yl}butanoic acid trifluoroacetate;
- 3-(3,5-Dimethoxyphenyl)-4-{3-[3-(5,6,7,8-tetrahydro-1,8-naphthyridin-2-yl)propyl]-1,2,4-oxadiazol-5-yl}butanoic acid trifluoroacetate;
- 80 3-(3,5-Dichlorophenyl)-4-{3-[3-(5,6,7,8-tetrahydro-1,8-naphthyridin-2-yl)propyl]-1,2,4-oxadiazol-5-yl}butanoic acid trifluoroacetate;
- 3-(3,5-Difluorophenyl)-4-{3-[3-(5,6,7,8-tetrahydro-1,8-naphthyridin-2-yl)propyl]-1,2,4-oxadiazol-5-yl}butanoic acid trifluoroacetate;
- 85 3-(3-Fluoro-4-methoxyphenyl)-4-{3-[3-(5,6,7,8-tetrahydro-1,8-naphthyridin-2-yl)propyl]-1,2,4-oxadiazol-5-yl}butanoic acid trifluoroacetate;
- 4-{3-[3-(5,6,7,8-Tetrahydro-1,8-naphthyridin-2-yl)propyl]-1,2,4-oxadiazol-5-yl}-3-[4-(trifluoromethyl)phenyl]butanoic acid trifluoroacetate;
- 3-(2-Methyl-1,3-thiazol-5-yl)-4-{3-[3-(5,6,7,8-tetrahydro-1,8-naphthyridin-2-yl)propyl]-1,2,4-oxadiazol-5-yl}butanoic acid trifluoroacetate;
- 90 3-(1-Phenyl-1H-pyrazol-4-yl)-4-{3-[3-(5,6,7,8-tetrahydro-1,8-naphthyridin-2-yl)propyl]-1,2,4-oxadiazol-5-yl}butanoic acid trifluoroacetate;
- 3-(1-Benzofuran-6-yl)-4-{3-[3-(5,6,7,8-tetrahydro-1,8-naphthyridin-2-yl)propyl]-1,2,4-oxadiazol-5-yl}butanoic acid hydrochloride;
- 95 3-(2,3-dihydro-1-benzofuran-6-yl)-4-{3-[3-(5,6,7,8-tetrahydro-1,8-naphthyridin-2-yl)propyl]-1,2,4-oxadiazol-5-yl}butanoic acid hydrochloride;
- 3-(1,3-Benzodioxol-5-yl)-4-{3-[3-[(pyridin-2-ylamino)methyl]phenyl]-1,2,4-oxadiazol-5-yl}butanoic acid hydrochloride;
- 3-(7-Fluoro-1,3-benzodioxol-5-yl)-4-{3-[3-(5,6,7,8-tetrahydro-1,8-naphthyridin-2-yl)propyl]-1,2,4-oxadiazol-5-yl}butanoic acid trifluoroacetate;
- 100

- 3-(1,3-Benzoxazol-6-yl)-4-{3-[3-(5,6,7,8-tetrahydro-1,8-naphthyridin-2-yl)propyl]-1,2,4-oxadiazol-5-yl}butanoic acid hydrochloride;
- 3-(3-Methyl-1,2,4-oxadiazol-5-yl)-4-{3-[3-(5,6,7,8-tetrahydro-1,8-naphthyridin-2-yl)propyl]-1,2,4-oxadiazol-5-yl}butanoic acid trifluoroacetate;
- 105 3-(3-Ethyl-1,2,4-oxadiazol-5-yl)-4-{3-[3-(5,6,7,8-tetrahydro-1,8-naphthyridin-2-yl)propyl]-1,2,4-oxadiazol-5-yl}butanoic acid trifluoroacetate;
- 3-(3-Phenyl-1,2,4-oxadiazol-5-yl)-4-{3-[3-(5,6,7,8-tetrahydro-1,8-naphthyridin-2-yl)propyl]-1,2,4-oxadiazol-5-yl}butanoic acid trifluoroacetate;
- [1-Benzoyl-4-({3-[3-(5,6,7,8-tetrahydro-1,8-naphthyridin-2-yl)propyl]-1,2,4-oxadiazol-5-yl)methyl}piperidin-4-yl]acetic acid trifluoroacetate;
- 110 [1-Benzoyl-4-({3-[4-(pyridin-2-ylamino)butyl]-1,2,4-oxadiazol-5-yl)methyl}piperidin-4-yl]acetic acid trifluoroacetate;
- [1-(tert-Butoxycarbonyl)-4-({3-[3-(5,6,7,8-tetrahydro-1,8-naphthyridin-2-yl)propyl]-1,2,4-oxadiazol-5-yl)methyl}piperidin-4-yl]acetic acid trifluoroacetate;
- 115 [1-(tert-Butoxycarbonyl)-4-({3-[4-(pyridin-2-ylamino)butyl]-1,2,4-oxadiazol-5-yl)methyl}piperidin-4-yl]acetic acid trifluoroacetate;
- 3-(4-Methylphenyl)-4-{3-[3-(5,6,7,8-tetrahydro-1,8-naphthyridin-2-yl)propyl]-1,2,4-oxadiazol-5-yl}butanoic acid trifluoroacetate;
- 3-(3-Chlorophenyl)-4-{3-[3-(5,6,7,8-tetrahydro-1,8-naphthyridin-2-yl)propyl]-1,2,4-oxadiazol-5-yl}butanoic acid hydrochloride;
- 120 3-(4-Methoxy-3-methylphenyl)-4-{3-[3-(5,6,7,8-tetrahydro-1,8-naphthyridin-2-yl)propyl]-1,2,4-oxadiazol-5-yl}butanoic acid hydrochloride;
- 3-[4-(Methylthio)phenyl]-4-{3-[3-(5,6,7,8-tetrahydro-1,8-naphthyridin-2-yl)propyl]-1,2,4-oxadiazol-5-yl}butanoic acid trifluoroacetate;
- 125 3-(1-Methyl-1H-indol-3-yl)-4-{3-[3-(5,6,7,8-tetrahydro-1,8-naphthyridin-2-yl)propyl]-1,2,4-oxadiazol-5-yl}butanoic acid trifluoroacetate;
- 3-(1,1'-Biphenyl-4-yl)-4-{3-[3-(5,6,7,8-tetrahydro-1,8-naphthyridin-2-yl)propyl]-1,2,4-oxadiazol-5-yl}butanoic acid hydrochloride;
- 3-(3-Bromophenyl)-4-{3-[3-(5,6,7,8-tetrahydro-1,8-naphthyridin-2-yl)propyl]-1,2,4-oxadiazol-5-yl}butanoic acid trifluoroacetate;
- 130 3-(4-Bromophenyl)-4-{3-[3-(5,6,7,8-tetrahydro-1,8-naphthyridin-2-yl)propyl]-1,2,4-oxadiazol-5-yl}butanoic acid hydrochloride;
- 3-(3-Phenoxyphenyl)-4-{3-[3-(5,6,7,8-tetrahydro-1,8-naphthyridin-2-yl)propyl]-1,2,4-oxadiazol-5-yl}butanoic acid trifluoroacetate;

- 135 3-[3-(Benzyloxy)phenyl]-4-{3-[3-(5,6,7,8-tetrahydro-1,8-naphthyridin-2-yl)propyl]-1,2,4-oxadiazol-5-yl}butanoic acid hydrochloride;
3-(3-Bromo-4-methoxyphenyl)-4-{3-[3-(5,6,7,8-tetrahydro-1,8-naphthyridin-2-yl)propyl]-1,2,4-oxadiazol-5-yl}butanoic acid trifluoroacetate;
4-{3-[3-(5,6,7,8-Tetrahydro-1,8-naphthyridin-2-yl)propyl]-1,2,4-oxadiazol-5-yl}-3-
- 140 (3,4,5-trimethoxyphenyl)butanoic acid trifluoroacetate;
3-(2-Naphthyl)-4-{3-[3-(5,6,7,8-tetrahydro-1,8-naphthyridin-2-yl)propyl]-1,2,4-oxadiazol-5-yl}butanoic acid hydrochloride;
3-(3-Nitrophenyl)-4-{3-[3-(5,6,7,8-tetrahydro-1,8-naphthyridin-2-yl)propyl]-1,2,4-oxadiazol-5-yl}butanoic acid hydrochloride;
- 145 3-(3-Methylphenyl)-4-{3-[3-(5,6,7,8-tetrahydro-1,8-naphthyridin-2-yl)propyl]-1,2,4-oxadiazol-5-yl}butanoic acid hydrochloride;
3-(2-Furyl)-4-{3-[3-(5,6,7,8-tetrahydro-1,8-naphthyridin-2-yl)propyl]-1,2,4-oxadiazol-5-yl}butanoic acid hydrochloride;
3-(2-Methylphenyl)-4-{3-[3-(5,6,7,8-tetrahydro-1,8-naphthyridin-2-yl)propyl]-1,2,4-
- 150 oxadiazol-5-yl}butanoic acid hydrochloride;
3-(1,3-benzodioxol-5-yl)-4-{3-[3-(3,4-dihydro-2H-pyrido[3,2-b][1,4]oxazin-6-yl)propyl]-1,2,4-oxadiazol-5-yl}butanoic acid, TFA;
4-{3-[3-(3,4-dihydro-2H-pyrido[3,2-b][1,4]oxazin-6-yl)propyl]-1,2,4-oxadiazol-5-yl}-3-(3,5-dimethoxyphenyl)butanoic acid, TFA;
- 155 3-Benzo[1,3]dioxol-5-yl-4-{3-[3-(6,7,8,9-tetrahydro-5-oxa-1,9-diaza-benzocyclohepten-2-yl)-propyl]-[1,2,4]oxadiazol-5-yl}-butyric acid;
3-(3-Fluoro-4-methoxyphenyl)-4-{3-[3-(6,7,8,9-tetrahydro-5-oxa-1,9-diaza-benzocyclohepten-2-yl)-propyl]-[1,2,4]oxadiazol-5-yl}-butyric acid;
3-(3,5-Difluorophenyl)-4-{3-[3-(6,7,8,9-tetrahydro-5-oxa-1,9-diaza-benzocyclohepten-
- 160 2-yl)-propyl]-[1,2,4]oxadiazol-5-yl}-butyric acid;
3-(3,5-Dimethoxyphenyl)-4-{3-[3-(6,7,8,9-tetrahydro-5-oxa-1,9-diaza-benzocyclohepten-2-yl)-propyl]-[1,2,4]oxadiazol-5-yl}-butyric acid;
3-(2-Methylbenzothiazol-5-yl)-4-{3-[3-(6,7,8,9-tetrahydro-5-oxa-1,9-diaza-benzocyclohepten-2-yl)-propyl]-[1,2,4]oxadiazol-5-yl}-butyric acid;
- 165 3-(1,3-benzodioxol-5-yl)-4-{3-[3-(1,2,3,5-tetrahydropyrido[2,3-e][1,4]oxazepin-8-yl)propyl]-1,2,4-oxadiazol-5-yl}butanoic acid, TFA;
3-(3,5-dimethoxyphenyl)-4-{3-[3-(1,2,3,5-tetrahydropyrido[2,3-e][1,4]oxazepin-8-yl)propyl]-1,2,4-oxadiazol-5-yl}butanoic acid, TFA;

- 170 3-(1,3-Benzodioxol-5-yl)-4-(3-{3-[6-(methylamino)pyridin-2-yl]propyl}-1,2,4-oxadiazol-5-yl)butanoic acid hydrochloride;
3-(3-Fluorophenyl)-4-(3-{3-[6-(methylamino)pyridin-2-yl]propyl}-1,2,4-oxadiazol-5-yl)butanoic acid trifluoroacetate;
3-(1,3-benzodioxol-5-yl)-4-(3-{3-[6-(ethylamino)pyridin-2-yl]propyl}-1,2,4-oxadiazol-5-yl)butanoic acid trifluoroacetate;
- 175 3-(3-Fuorophenyl)-4-(3-{3-[6-(methylamino) pyridin-2- yl]propyl}-1,2,4-oxadiazol-5-yl)butanoic acid trifluoroacetate;
3-(1,3-Benzodioxol-5-yl)-4-(3-{4-[(4-methylpyridin-2-yl)amino]butyl}-1,2,4-oxadiazol-5-yl)butanoic acid;
3-(1,3-benzodioxol-5-yl)-4-(3-{4-[(6-methylpyridin-2-yl)amino]butyl}-1,2,4-oxadiazol-
- 180 5-yl)butanoic acid;
(2-{6-[2-(5,6,7,8-Tetrahydro-1,8-naphthyridin-2-yl)ethoxy]pyridin-3-yl}cyclopropyl)acetic acid;
3-Methyl-4-{6-[2-(5,6,7,8-tetrahydro-1,8-naphthyridin-2-yl)ethoxy]pyridin-3-yl}butanoic acid;
- 185 3-(1,3-benzodioxol-5-yl)-4-{5-[3-(5,6,7,8-tetrahydro-1,8-naphthyridin-2-yl)propyl]-1,3,4-oxadiazol-2-yl}butanoic acid trifluoroacetate;
3-(3-fluorophenyl)-4-{5-[3-(5,6,7,8-tetrahydro-1,8-naphthyridin-2-yl)propyl]-1,3,4-oxadiazol-2-yl}butanoic acid trifluoroacetate;
3-(3-Fluoro-4-methoxyphenyl)-4-{5-[3-(5,6,7,8-tetrahydro-1,8-naphthyridin-2-
- 190 yl)propyl]-1,3,4-oxadiazol-2-yl}butanoic acid trifluoroacetate;
3-(3,5-Dimethoxyphenyl)-4-{5-[3-(5,6,7,8-tetrahydro-1,8-naphthyridin-2-yl)propyl]-1,3,4-oxadiazol-2-yl}butanoic acid trifluoroacetate;
3-(2-Methyl-1,3-thiazol-5-yl)-4-{5-[3-(5,6,7,8-tetrahydro-1,8-naphthyridin-2-yl)propyl]-1,3,4-oxadiazol-2-yl}butanoic acid trifluoroacetate;
- 195 3-(4-Fluorophenyl)-4-{5-[3-(5,6,7,8-tetrahydro-1,8-naphthyridin-2-yl)propyl]-1,3,4-oxadiazol-2-yl}butanoic acid trifluoroacetate;
3-(3,5-Difluorophenyl)-4-{5-[3-(5,6,7,8-tetrahydro-1,8-naphthyridin-2-yl)propyl]-1,3,4-oxadiazol-2-yl}butanoic acid trifluoroacetate;
- 200 3-(3,5-Difluorophenyl)-4-{5-[3-(5,6,7,8-tetrahydro-1,8-naphthyridin-2-yl)propyl]-1,3,4-thiadiazol-2-yl}butanoic acid trifluoroacetate;
3-(4-Fluorophenyl)-4-{5-[3-(5,6,7,8-tetrahydro-1,8-naphthyridin-2-yl)propyl]-1,3,4-thiadiazol-2-yl}butanoic acid trifluoroacetate;

- 3-(2-Methyl-1,3-thiazol-5-yl)-4-{5-[3-(5,6,7,8-tetrahydro-1,8-naphthyridin-2-yl)propyl]-1,3,4-thiadiazol-2-yl}butanoic acid trifluoroacetate;
- 205 3-(1,3-Benzodioxol-5-yl)-4-{5-[3-(5,6,7,8-tetrahydro-1,8-naphthyridin-2-yl)propyl]-1,3,4-thiadiazol-2-yl}butanoic acid trifluoroacetate;
- 3-(1,3-benzodioxol-5-yl)-4-{3-[2-(5,6,7,8-tetrahydro-1,8-naphthyridin-2-yl)ethoxy]isoxazol-5-yl}butanoic acid;
- 210 3-(1,3-benzodioxol-5-yl)-4-{5-[3-(5,6,7,8-tetrahydro-1,8-naphthyridin-2-yl)propyl]-2H-tetraazol-2-yl}butanoic acid;
- 3-(1,3-benzodioxol-5-yl)-4-{5-[3-(5,6,7,8-tetrahydro-1,8-naphthyridin-2-yl)propyl]-1H-tetraazol-1-yl}butanoic acid;
- 3-(1,3-benzodioxol-5-yl)-4-{3-[2-(5,6,7,8-tetrahydro-1,8-naphthyridin-2-yl)ethoxy]-1H-pyrazol-5-yl}butanoic acid;
- 215 3-(1,3-benzodioxol-5-yl)-4-{3-[3-(4,5-dihydro-1H-imidazol-2-ylamino)propoxy]isoxazol-5-yl}butanoic acid;
- 3-[2-(4-chlorophenyl)-1,3-thiazol-5-yl]-4-{3-[2-(5,6,7,8-tetrahydro-1,8-naphthyridin-2-yl)ethoxy]isoxazol-5-yl}butanoic acid;
- 3-Benzo[1,3]dioxol-5-yl-4-{3-[2-(6,7,8,9-tetrahydro-5-oxa-1,9-diazabenzocyclohepten-2-yl)-ethoxy]-isoxazol-5-yl}-butyric acid;
- 220 3-Benzo[1,3]dioxol-5-yl-4-{3-oxo-2-[2-(6,7,8,9-tetrahydro-5-oxa-1,9-diazabenzocyclohepten-2-yl)-ethyl]-2,3-dihydro-isoxazol-5-yl}-butyric acid;
- 3-(1,3-benzodioxol-5-yl)-4-{3-[2-(3,4-dihydro-2H-pyrido[3,2-b][1,4]oxazin-6-yl)ethoxy]isoxazol-5-yl}butanoic acid, TFA;
- 225 3-(1,3-benzodioxol-5-yl)-4-{2-[2-(3,4-dihydro-2H-pyrido[3,2-b][1,4]oxazin-6-yl)ethyl]-3-oxo-2,3-dihydroisoxazol-5-yl}butanoic acid, TFA;
- 3-(1,3-benzodioxol-5-yl)-4-{3-[2-(1,2,3,5-tetrahydropyrido[2,3-e][1,4]oxazepin-8-yl)ethoxy]isoxazol-5-yl}butanoic acid, TFA;
- 3-(1,3-benzodioxol-5-yl)-4-{3-oxo-2-[2-(1,2,3,5-tetrahydropyrido[2,3-e][1,4]oxazepin-8-yl)ethyl]-2,3-dihydroisoxazol-5-yl}butanoic acid, TFA;
- 230 3-(1,3-benzodioxol-5-yl)-4-{3-[2-[5-(methoxymethyl)-6-(methylamino)pyridin-2-yl]ethoxy]isoxazol-5-yl}butanoic acid, TFA;
- 3-(1,3-Benzodioxol-5-yl)-4-{3-[3-(5,5-dimethyl-5,6,7,8-tetrahydro-1,8-naphthyridin-2-yl)propyl]-1,2,4-oxadiazol-5-yl}butanoic acid;
- 235 3-(1,3-benzodioxol-5-yl)-4-{3-[3-(1-methyl-1,2,3,4-tetrahydropyrido[2,3-b]pyrazin-6-yl)propyl]-1,2,4-oxadiazol-5-yl}butanoic acid;

3-(2-methyl-1,3-benzothiazol-5-yl)-4-{3-[3-(1-methyl-1,2,3,4-tetrahydropyrido[2,3-b]pyrazin-6-yl)propyl]-1,2,4-oxadiazol-5-yl}butanoic acid;
 3-(3-fluoro-4-methoxyphenyl)-4-{3-[3-(1-methyl-1,2,3,4-tetrahydropyrido[2,3-b]pyrazin-6-yl)propyl]-1,2,4-oxadiazol-5-yl}butanoic acid;
 240 3-(6-methoxypyridin-3-yl)-4-{3-[3-(1-methyl-1,2,3,4-tetrahydropyrido[2,3-b]pyrazin-6-yl)propyl]-1,2,4-oxadiazol-5-yl}butanoic acid;
 3-(1,3-benzodioxol-5-yl)-4-(3-{[2-(5,6,7,8-tetrahydro-1,8-naphthyridin-2-yl)ethyl]thio}-1H-1,2,4-triazol-5-yl)butanoic acid;
 245 3-(1,3-benzodioxol-5-yl)-4-(1-methyl-5-{[2-(5,6,7,8-tetrahydro-1,8-naphthyridin-2-yl)ethyl]thio}-1H-1,2,4-triazol-3-yl)butanoic acid;
 3-(1,3-benzodioxol-5-yl)-4-(4-methyl-5-{[2-(5,6,7,8-tetrahydro-1,8-naphthyridin-2-yl)ethyl]thio}-4H-1,2,4-triazol-3-yl)butanoic acid;
 3-(1,3-benzodioxol-5-yl)-4-{3-[2-(1-methyl-1,2,3,4-tetrahydropyrido[2,3-b]pyrazin-6-yl)ethoxy]isoxazol-5-yl}butanoic acid;
 250 3-(1,3-benzodioxol-5-yl)-4-(3-[2-[6-(methylamino)pyridin-2-yl]ethoxy]isoxazol-5-yl)butanoic acid; and
 3-(6-methoxypyridin-3-yl)-4-{3-[2-(5,6,7,8-tetrahydro-1,8-naphthyridin-2-yl)ethoxy]isoxazol-5-yl}butanoic acid.

2. A method for the treatment or prevention of conditions mediated by the $\alpha_v\beta_3$ integrin in a mammal in need of such treatment, the method comprising administering to the subject a therapeutically effective amount of a composition of Claim 1.

3. The method according to Claim 2 wherein the condition treated is selected from the group consisting of tumor metastasis, solid tumor growth, angiogenesis, osteoporosis, humoral hypercalcemia of malignancy, smooth muscle cell migration, restenosis, atherosclerosis, macular degeneration, retinopathy, and
 5 arthritis.

4. A method for the treatment or prevention of conditions mediated by the $\alpha_v\beta_5$ integrin in a mammal in need of such treatment, the method comprising

administering to the subject a therapeutically effective amount of a composition of Claim 1.

5. The method according to Claim 4 wherein the condition treated is selected from the group consisting of tumor metastasis, solid tumor growth, angiogenesis, osteoporosis, humoral hypercalcemia of malignancy, smooth muscle cell migration, restenosis, atherosclerosis, macular degeneration, retinopathy, and arthritis.